

Please cancel claims <sup>/ / / / / / / / / /</sup> 19, 22, 28, 29, 30, 35, 36, 37, 42, 43, 44, 49, 50, and 51, without prejudice.

<sup>/ / / / /</sup> Claims 15, 21, 23, 24, 25, 32, 39, and 46 have been amended as follows.

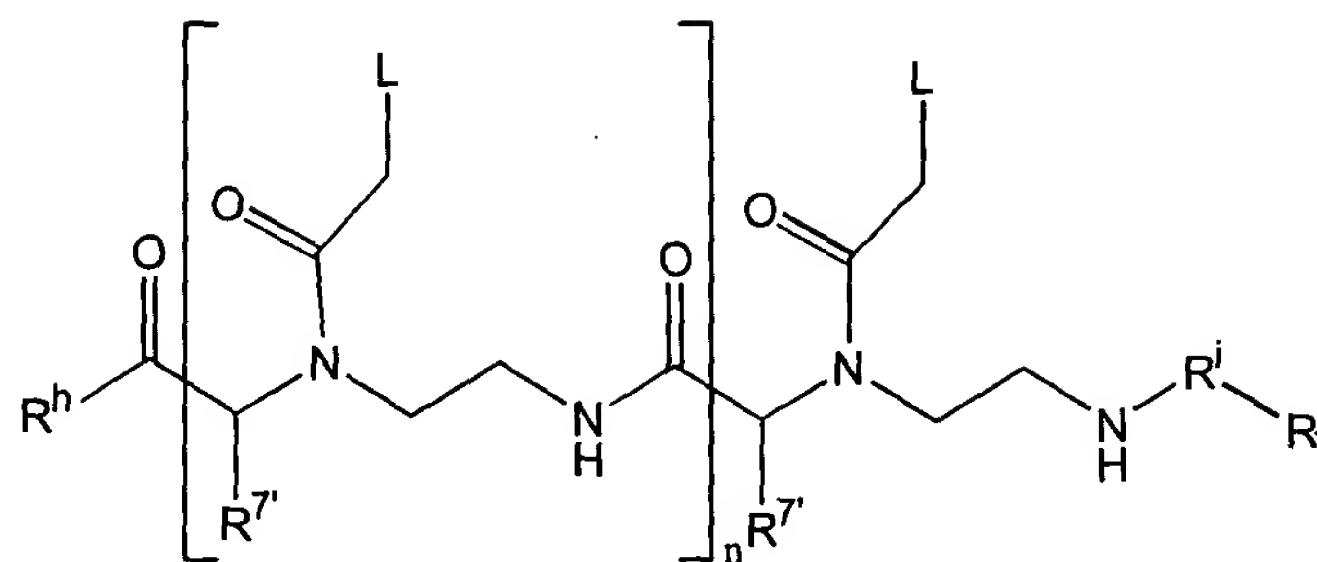
15. A method of modulating cellular uptake and distribution of a peptide nucleic acid comprising the steps of:

- (a) derivatizing a backbone position of said peptide nucleic acid; and
- (b) conjugating the derivatized peptide nucleic acid of step (a) with a group selected from alkyl, lipid, and steroid.

<sup>SUB E</sup> 21. A method of modulating cellular uptake and distribution of a peptide nucleic acid comprising the steps of:

- (a) conjugating said peptide nucleic acid with a group selected from alkyl, lipid, and steroid; and
- (b) introducing the conjugated peptide nucleic acid of step (a) into liposomes.

<sup>D</sup> 23. A composition comprising a peptide nucleic acid having formula:



wherein:

each L is, independently, a naturally-occurring nucleobase or a non-naturally-occurring nucleobase;

each R<sup>7</sup> is hydrogen or the side chain of a naturally-occurring or non-naturally-occurring amino acid, at least one R<sup>7</sup> being the side chain of a naturally-occurring or non-naturally-occurring amino acid;

R<sup>h</sup> is OH, NH<sub>2</sub>, or NHLysNH<sub>2</sub>;

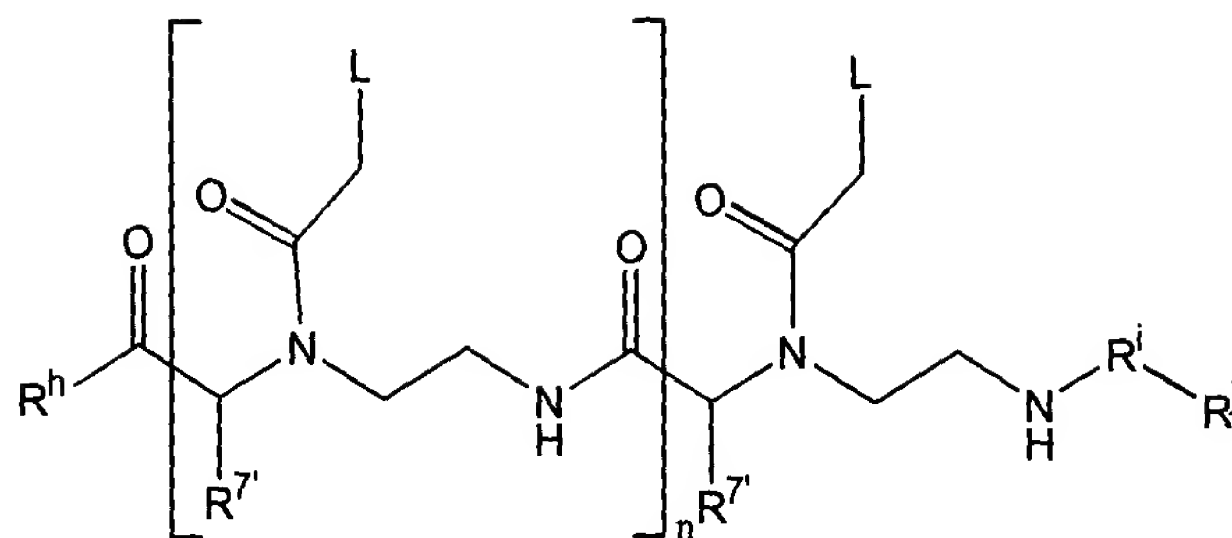
each of R<sup>i</sup> and R<sup>j</sup> is, independently, a group selected from alkyl, lipid, and steroid; or R<sup>i</sup> and R<sup>j</sup>, together, are a group selected from alkyl, lipid and steroid; and

n is an integer from 1 to 30;

and at least one pharmaceutically acceptable carrier, binder, thickener, diluent, buffer, preservative or surface active agent.

24. A composition comprising a peptide nucleic acid incorporated into a liposome, said peptide nucleic acid having formula:

3  
continued



wherein:

each L is, independently, a naturally-occurring nucleobase or a non-naturally-occurring nucleobase;

each R<sup>7</sup> is hydrogen or the side chain of a naturally-occurring or non-naturally-occurring amino acid;

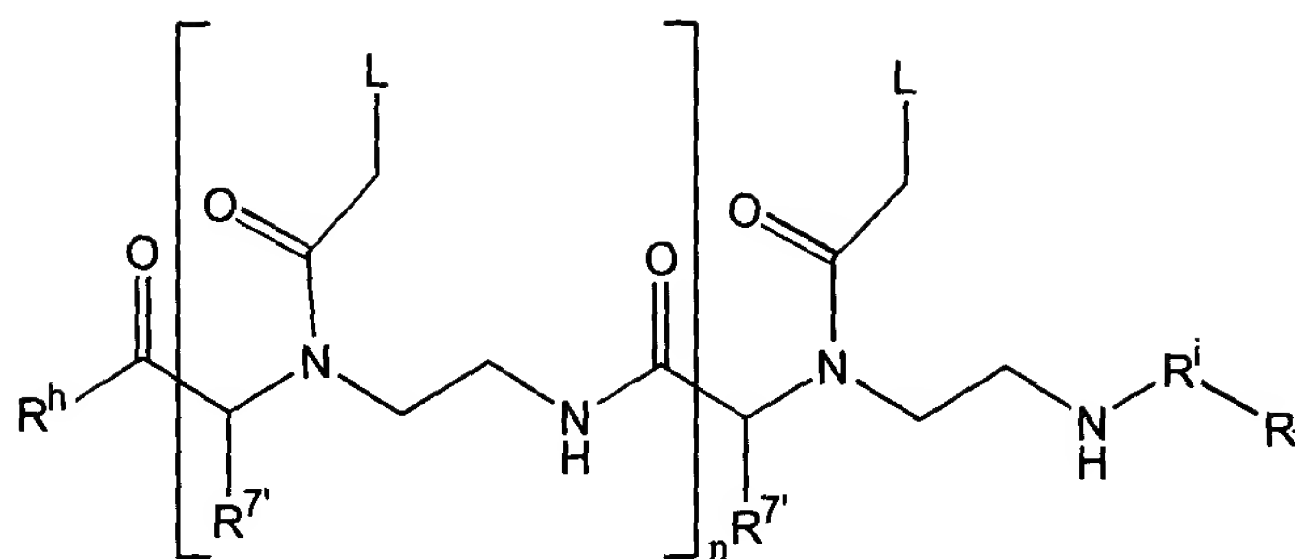
R<sup>h</sup> is OH, NH<sub>2</sub>, or NHLysNH<sub>2</sub>;

each of R<sup>i</sup> and R<sup>j</sup> is, independently, a group selected from alkyl, lipid, and steroid; or R<sup>i</sup> and R<sup>j</sup>, together, are a group selected from alkyl, lipid and steroid; and

n is an integer from 1 to 30;

and at least one pharmaceutically acceptable carrier, binder, thickener, diluent, buffer, preservative or surface active agent.

25. A method of modulating cellular uptake and distribution of a peptide nucleic acid in a cell or tissue comprising administering to the cell or tissue a peptide nucleic acid having formula:



wherein:

each L is, independently, a naturally-occurring nucleobase or a non-naturally-occurring nucleobase;

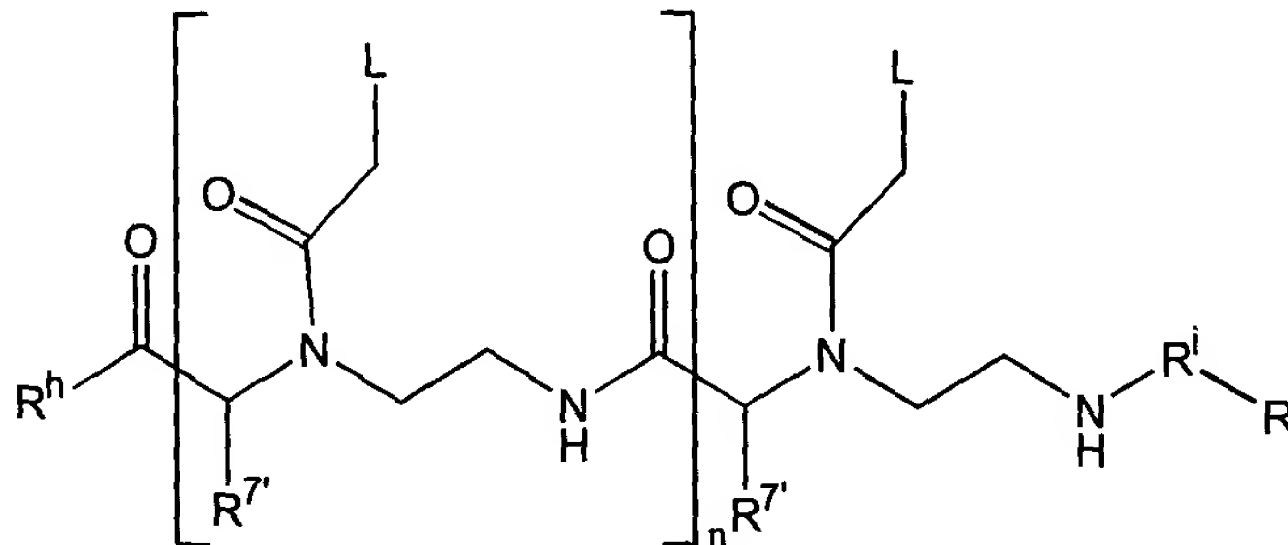
each R<sup>7'</sup> is hydrogen or the side chain of a naturally-occurring or non-naturally-occurring amino acid, at least one R<sup>7'</sup> being the side chain of a naturally-occurring or non-naturally-occurring amino acid;

R<sup>h</sup> is OH, NH<sub>2</sub>, or NHLysNH<sub>2</sub>;

each of R<sup>i</sup> and R<sup>j</sup> is, independently, a group selected from alkyl, lipid, and steroid; or R<sup>i</sup> and R<sup>j</sup>, together, are a group selected from alkyl, lipid, and steroid; and

n is an integer from 1 to 30.

32. A method of modulating cellular uptake and distribution of a peptide nucleic acid in a cell or tissue comprising administering to the cell or tissue a composition comprising a peptide nucleic acid incorporated into a liposome, said peptide nucleic acid having formula:



wherein:

each L is, independently, a naturally-occurring nucleobase or a non-naturally-occurring nucleobase;

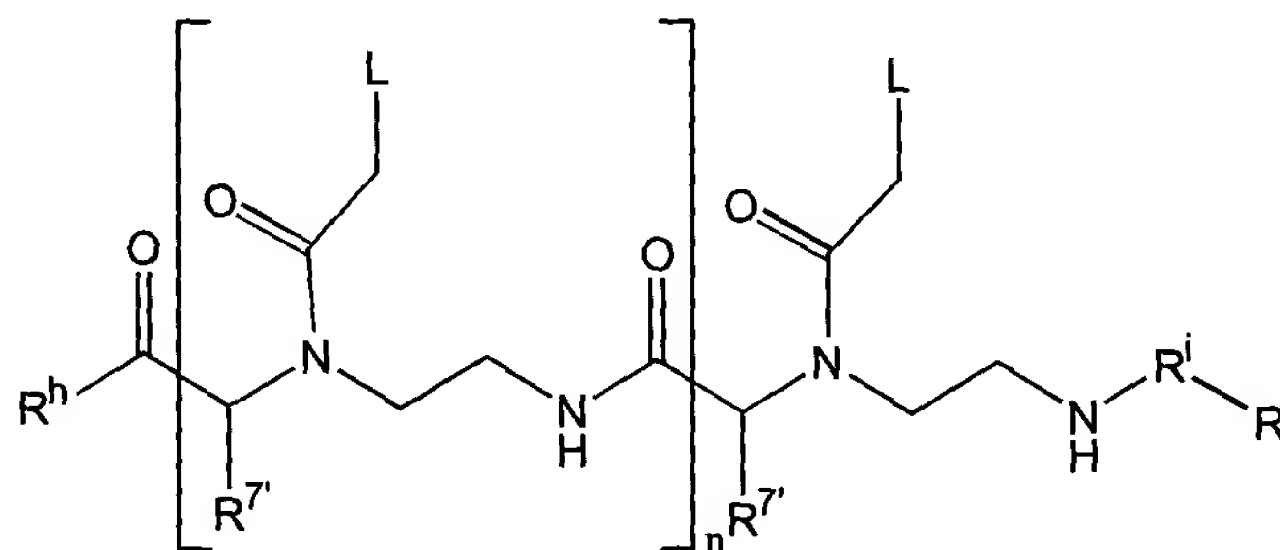
each R<sup>7</sup> is hydrogen or the side chain of a naturally-occurring or non-naturally-occurring amino acid;

R<sup>h</sup> is OH, NH<sub>2</sub>, or NHLysNH<sub>2</sub>;

each of R<sup>i</sup> and R<sup>j</sup> is, independently, a group selected from alkyl, lipid, and steroid; or R<sup>i</sup> and R<sup>j</sup>, together, are a group selected from alkyl, lipid, and steroid; and

n is an integer from 1 to 30.

39. A method of modulating gene expression in an animal comprising administering to the animal a therapeutically effective amount of a peptide nucleic acid of formula:



wherein:

each L is, independently, a naturally-occurring nucleobase or a non-naturally-occurring nucleobase;

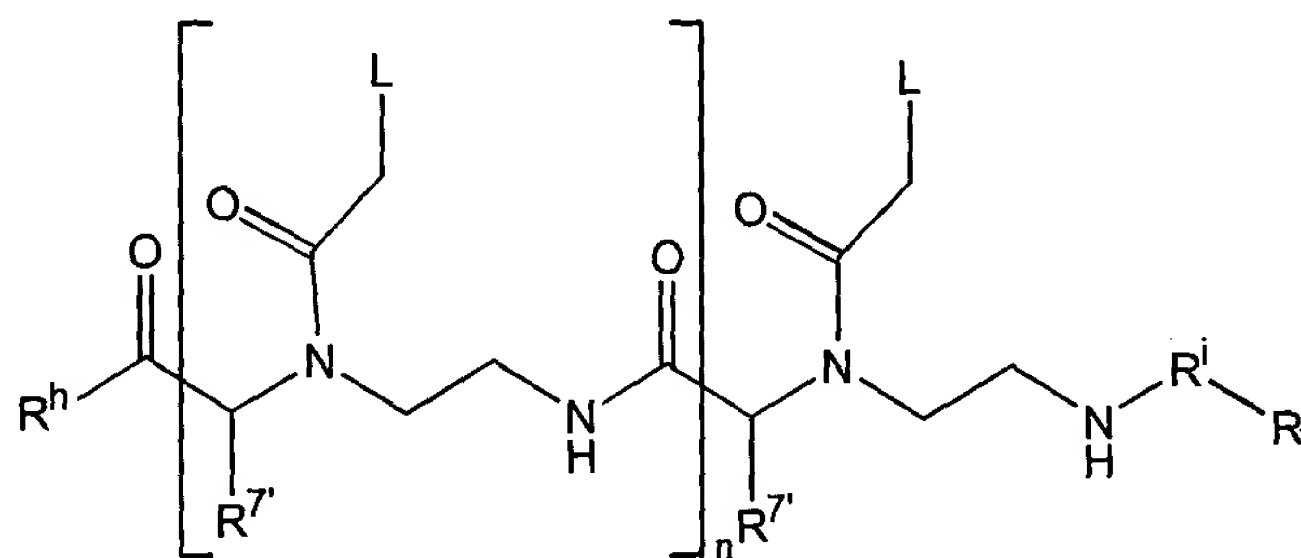
each R<sup>7</sup> is hydrogen or the side chain of a naturally-occurring or non-naturally-occurring amino acid, at least one R<sup>7</sup> being the side chain of a naturally-occurring or non-naturally-occurring amino acid;

R<sup>h</sup> is OH, NH<sub>2</sub>, or NHLysNH<sub>2</sub>;

*D<sup>5</sup> control*

each of  $R^i$  and  $R^j$  is, independently, a group selected from alkyl, lipid, and steroid; or  $R^i$  and  $R^j$ , together, are a group selected from alkyl, lipid, and steroid; and  
 $n$  is an integer from 1 to 30.

46. A method of modulating gene expression in an animal comprising administering to the animal a therapeutically effective amount of a composition comprising a peptide nucleic acid incorporated into a liposome, said peptide nucleic acid having formula:



wherein:

each  $L$  is, independently, a naturally-occurring nucleobase or a non-naturally-occurring nucleobase;

each  $R^7$  is hydrogen or the side chain of a naturally-occurring or non-naturally-occurring amino acid;

$R^h$  is  $\text{OH}$ ,  $\text{NH}_2$ , or  $\text{NHLysNH}_2$ ;

each of  $R^i$  and  $R^j$  is, independently, a group selected from alkyl, lipid, and steroid; or  $R^i$  and  $R^j$ , together, are a group selected from alkyl, lipid, and steroid; and

$n$  is an integer from 1 to 30.

# REMARKS

Claims 15-52 are pending in this application. Claims 15, 21, 23, 24, 25, 32, 39, and 46